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U.S. National Phase of PCT/AU00/00189 New Claims and Redlined Version of Amended Claims

- 1. (Amended) An isolated and purified nucleic acid sequence comprising a polynucleotide sequence encoding a polypeptide of an antibody-(_or fragment thereof), wherein said antibody-(_or fragment thereof)—has binding affinity to a p53 protein or a portion thereof in vertebrates, and wherein said nucleic acid sequence is obtained from a vertebrate host expressing an immune response against a naturally-occurring disease.
- 2. (Amended) A-<u>The</u> nucleic acid sequence according to claim 1, wherein said immune response is <u>characterised</u> characterized by expression of at least one p53 antibody.
- 3. (Amended) A-The nucleic acid sequence according to claim 1 or claim 2 comprising a polynucleotide sequence encoding an F_{ab} antibody fragment (, or fragment thereof,) having binding affinity to a p53 protein or a portion thereof in vertebrates.
- 4. (Amended) An isolated and purified nucleic acid sequence encoding a polypeptide of an antibody-(, or fragment thereof,) comprising a polynucleotide sequence selected from the group consisting of SEQ ID Nos 1-30, wherein said antibody-(, or fragment thereof,) has binding affinity to a p53 protein or a portion thereof.
- 5. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u>-1 to 4, wherein the nucleic acid sequence is DNA.
- 6. (Amended) A-<u>The</u> nucleic acid sequence according to any one of claims claim 1 to 4, wherein the nucleic acid sequence is RNA.
- 7. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to 6, wherein the nucleic acid sequence comprises a polynucleotide sequence(s) or sequences, or an analogue thereof, encoding an antibody fragment or other immunologically active fragment thereof, -wherein the antibody-(_or fragment thereof_) has binding affinity to a p53 protein or a portion thereof in vertebrates.
- 8. (Amended) A-The nucleic acid sequence according to claim 7, wherein the antibody fragment or other immunologically active fragment comprises at least one complementarity determining region.

- 9. (Amended) A-<u>The</u> nucleic acid sequence according to claim 7-or claim 8, wherein the antibody fragment comprises at least one functional antigen-binding domain.
- 10. (Amended) A-The nucleic acid sequence according to any one of claims claim 7 to 9, wherein the antibody fragment is selected from the group consisting of: Fv, Fab, F(ab)2, scFv (single chain Fv), dAb (single domain antibody), bi-specific antibodies, diabodies and triabodies.
- 11. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to 10, wherein the antibody-(,_or fragment thereof,) has binding affinity for residues of one or more of the N-terminus, the C-terminus or the central domain of a p53 protein or a portion thereof.
- 12. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to-11, wherein the antibody-(_or fragment thereof_) has binding affinity for residues of the N-terminus of a p53 protein or a portion thereof.
- 13. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1 to 12, wherein the antibody-(_or fragment thereof_) has binding affinity for residues about 10 to about 55 of the N-terminus of a p53 protein or portion thereof.
- 14. (Amended) A-The nucleic acid sequence according to any one of claimsclaim 1-to 12, wherein the antibody-1_or fragment thereof_) has binding affinity for residues about 10 to about 25 of the N-terminus of a p53 protein or portion thereof.
- 15. (Amended) A-The nucleic acid sequence according to any one of claimsclaim 1-to 12, wherein the antibody-(_or fragment thereof_) has binding affinity for residues about 40 to about 50 of the N-terminus of a p53 protein or portion thereof.
- 16. (Amended) A-The nucleic acid sequence according to any one of claimsclaim 1-to-12, wherein the antibody-(_or fragment thereof_) has binding affinity for residues about 27 to about 44 of the N-terminus of a p53 protein or portion thereof.
- 17. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to-12, wherein the antibody-(_or fragment thereof)-_has binding affinity for residues about 40 to about 44 of the N-terminus of a p53 protein or portion thereof.

- 18. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to-11, wherein the antibody-1_or fragment thereof)-_has binding affinity for residues of the central domain of a p53 protein or a portion thereof.
- 19. (Amended) A-The nucleic acid sequence according to any one of claimsclaim 1-to 18, wherein said sequence comprises a polynucleotide sequence encoding a polypeptide of an antibody-(_, or fragment thereof)-_, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said polynucleotide sequence encodes an immunoglobulin light chain variable region polypeptide or an immunoglobulin heavy chain variable region polypeptide.
- 20. (Amended) A-The nucleic acid sequence according to any one of claimsclaim 1 to 19, wherein said sequence comprises a polynucleotide sequence encoding a polypeptide of an antibody-(_or fragment thereof)-_having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said nucleic acid sequence comprises a first polynucleotide sequence encoding an immunoglobulin light chain variable region polypeptide, and a second polynucleotide sequence encoding an immunoglobulin heavy chain variable region polypeptide.
- 21. (Amended) A-<u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1-to 20, wherein the vertebrate is selected from the group consisting of human, non-human primate, murine, bovine, ovine, equine, caprine, leporine, avian, feline and canine.
- 22. (Amended) A <u>The</u> nucleic acid sequence according to <u>any one of claimsclaim</u> 1 to 21, wherein the vertebrate is human.
- 23. (Amended) An isolated and purified nucleic acid sequence comprising an analogue of the nucleic acid sequence according to any one of claimsclaim 1-to 22, wherein said analogue encodes a polypeptide having a biological activity which is functionally the same as the polypeptide(s) encoded by said polynucleotide sequence.
- 24. (Amended) A-The nucleic acid sequence according to any one of claims claim 1 to 23, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.

- 25. (Amended) A-<u>The</u> nucleic acid sequence according to claim 24, wherein the disease is cancer.
- 26. (Amended) A-The nucleic acid sequence according to claim 25, wherein the cancer is selected from the group consisting of carcinogenic tumourtumors; tumourtumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumourtumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumourtumors, such as sarcoma; and haemopoietic tumourtumors, such as B cell lymphoma.
- 27. (Amended) A polypeptide of an antibody-{_or fragment thereof}-_having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said polypeptide is obtained from a vertebrate host expressing an immune response against a naturally-occurring disease.
- 28. (Amended) A-<u>The</u> polypeptide according to claim 27, wherein said immune response is characterisedcharacterized by expression of at least one p53 antibody.
- 29. (Amended) An isolated and purified polypeptide, wherein said polypeptide is encoded by the nucleic acid sequence according to any one of claimsclaim 1-to-26.
- 30. (Amended) An isolated and purified polypeptide of an antibody—(_or fragment thereof)—, comprising an amino acid sequence selected from the group consisting of SEQ ID Nos 31-60, wherein said antibody—(_or fragment thereof)—has binding affinity to a p53 protein or a portion thereof.
- 31. (Amended) A-The polypeptide according to any one of claimsclaim 27-to 30, wherein said polypeptide is selected from the group consisting of: Fv, F_{ab}, F(ab)₂, scFv (single chain Fv), dAb (single domain antibody), bi-specific antibodies, diabodies and triabodies.
- 32. (Amended) A-The polypeptide according to any one of claims claim 27 to 31, wherein said polypeptide has binding affinity to a p53 protein or a portion thereof.

- 33. (Amended) A-<u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27-to 32, wherein said polypeptide has binding affinity for residues of one or more of the N-terminus, the C-terminus or the central domain of a p53 protein or a portion thereof.
- 34. (Amended) A-<u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27-to-33, wherein said polypeptide has binding affinity for residues of the N-terminus of a p53 protein or a portion thereof.
- 35. (Amended) A <u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27 to 34, wherein said polypeptide has binding affinity for residues about 10 to about 55 of the N-terminus of a p53 protein or portion thereof.
- 36. (Amended) A-The polypeptide according to any one of claimsclaim 27-to-34, wherein said polypeptide has binding affinity for residues about 10 to about 25 of the N-terminus of a p53 protein or portion thereof.
- 37. (Amended) A-<u>The polypeptide according to any one of claimsclaim</u> 27-to-34, wherein said polypeptide has binding affinity for residues about 40 to about 50 of the N-terminus of a p53 protein or portion thereof.
- 38. (Amended) A-<u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27-to 34, wherein said polypeptide has binding affinity for residues about 27 to about 44 of the N-terminus of a p53 protein or portion thereof.
- 39. (Amended) A-The polypeptide according to any one of claimsclaim 27-to-34, wherein said polypeptide has binding affinity for residues about 40 to about 44 of the N-terminus of a p53 protein or portion thereof.
- 40. (Amended) A-<u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27-to 33, wherein said polypeptide has binding affinity for residues of the central domain of a p53 protein or a portion thereof.
- 41. (Amended) An isolated and purified polypeptide, wherein said polypeptide is a homologous polypeptide of the polypeptide according to any one of claims 27-to 40.

- 42. (Amended) A-The polypeptide according to claim 41, wherein said polypeptide is at least 45% homologous to the a polypeptide according to any one of claims 27 to 40of an antibody, or fragment thereof, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said polypeptide of an antibody is obtained from a vertebrate host expressing an immune response against a naturally-occurring disease.
- 43. (Amended) A-The polypeptide according to claim 41, wherein said polypeptide is at least 75% homologous to the polypeptide of an antibody, or fragment thereof, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said polypeptide of an antibody is obtained from a vertebrate host expressing an immune response against a naturally-occurring diseaseaccording to any one of claims 27 to 40.
- 44. (Amended) A-The polypeptide according to claim 41, wherein said polypeptide is at least 95-99% homologous to the polypeptide of an antibody, or fragment thereof, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said polypeptide of an antibody is obtained from a vertebrate host expressing an immune response against a naturally-occurring disease according to any one of claims 27 to 40.
- 45. (Amended) A-<u>The</u> polypeptide according to <u>any one of claimsclaim</u> 27-to 44, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 46. (Amended) A-The polypeptide according to claim 45, wherein the disease is cancer.
- 47. (Amended) A-The polypeptide according to claim 46, wherein the cancer is selected from the group consisting of carcinogenic tumourtumors; tumourtumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumourtumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumourtumors, such as sarcoma; and haemopoietic tumourtumors, such as B cell lymphoma.
- 49. (Amended) A-The peptide fragment according to claim 48, wherein said peptide fragment comprises between about 5 and about 50 contiguous amino acids of any one of SEQ ID Nos 31-60.

- 50. (Amended) A-The peptide fragment according to any one of claimsclaim 48 to 49, wherein said peptide fragment comprises between about 5 and about 30 contiguous amino acids of any one of SEQ ID Nos 31-60.
- 51. (Amended) A-The peptide fragment according to any one of claimsclaim 48 to 50, wherein said peptide fragment comprises between about 8 and about 20 contiguous amino acids of any one of SEQ ID Nos 31-60.
- 52. (Amended) A-<u>The</u> peptide fragment according to claim 48, wherein said peptide fragment is derived from the complementarity determining region.
- 53. (Amended) A-<u>The</u> peptide fragment according to any one of claims claim 48 to 52, wherein said immune response is an idiotypic response.
- 54. (Amended) A-<u>The</u> peptide fragment according to any one of claims claim 48 to 53, wherein the vertebrate is human.
- 56. (Amended) An The antibody or fragment thereof according to claim 55, wherein said immune response is characterised characterized by expression of a p53 antibody.
- 57. (Amended) An-The antibody, or fragment thereof, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said antibody or fragment thereof is comprised of the polypeptide according to any one of claimsclaim 27 to 47.
- 58. (Amended) An-The antibody, or fragment thereof, having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said antibody or fragment thereof is encoded by the nucleic acid sequence according to any one of claimsclaim 1 to 26.
- 59. (Amended) An The antibody fragment according to any one of claims claim 55 to 58, wherein said fragment is an immunologically active fragment.
- 60. (Amended) An The antibody fragment according to any one of claims claim 55 to 59, wherein said fragment comprises at least one complementarity determining region.
- 61. (Amended) An-The antibody fragment according to any one of claims claim 55-to 60, wherein said fragment is selected from the group consisting of: Fv, Fab, F(ab)2, scFv

(single chain Fv), dAb (single domain antibody), bi-specific antibodies, diabodies and triabodies.

- 62. (Amended) An <u>The antibody</u>, or fragment thereof, according to <u>any one of claimsclaim</u> 55 to 61, which is a polyclonal antibody.
- 63. (Amended) An <u>The antibody</u>, or fragment thereof, according to <u>any one of claimsclaim</u> 55 to 61, which is a monoclonal antibody.
- 64. (Amended) An-<u>The</u> antibody or fragment thereof according to <u>any one of claimsclaim</u> 57 to 63, wherein said antibody or fragment thereof has binding affinity for residues of one or more of the N-terminus, the C-terminus or the central domain of a p53 protein or a portion thereof.
- 65. (Amended) An-The antibody or fragment thereof according to any one of claimsclaim 57 to 64, wherein said antibody or fragment thereof has binding affinity for residues of the N-terminus of a p53 protein or a portion thereof.
- 66. (Amended) An-<u>The</u> antibody or fragment thereof according to any one of claimsclaim 57 to 65, wherein said antibody or fragment thereof has binding affinity for residues about 10 to about 55 of the N-terminus of a p53 protein or portion thereof.
- 67. (Amended) An-The antibody or fragment thereof according to any one of claims claim 57 to 65, wherein said antibody or fragment thereof has binding affinity for residues about 10 to about 25 of the N-terminus of a p53 protein or portion thereof.
- 68. (Amended) An-The antibody or fragment thereof according to any one of claimsclaim 57 to 65, wherein said antibody or fragment thereof has binding affinity for residues about 40 to about 50 of the N-terminus of a p53 protein or portion thereof.
- 69. (Amended) An-<u>The</u> antibody or fragment thereof according to any one of claimsclaim 57 to 65, wherein said antibody or fragment thereof has binding affinity for residues about 27 to about 44 of the N-terminus of a p53 protein or portion thereof.
- 70. (Amended) An-The antibody or fragment thereof according to any one of claims claim 57 to 65, wherein said antibody or fragment thereof has binding affinity for residues about 40 to about 44 of the N-terminus of a p53 protein or portion thereof.

- 71. (Amended) An-<u>The</u> antibody or fragment thereof according to any one of claimsclaim 57 to 64, wherein said antibody or fragment thereof has binding affinity for residues of the central domain of a p53 protein or a portion thereof.
- 72. (Amended) An-The antibody or fragment thereof according to any one of claimsclaim 55 to 71, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 73. (Amended) An-The antibody or fragment thereof according to claim 72, wherein the disease is cancer.
- 74. (Amended) An-The antibody or fragment thereof according to claim 73, wherein the cancer is selected from the group consisting of carcinogenic tumourtumors; tumourtumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumourtumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumourtumors, such as sarcoma; and haemopoietic tumourtumors, such as B cell lymphoma.
- 75. (Amended) A vector comprising the nucleic acid sequence according to any one of claimsclaim 1-to-26.
- 76. (Amended) A-<u>The</u> vector according to claim 75, wherein said vector is selected from the group consisting of viral, plasmid, bacteriophage, phagemid, cosmid, bacterial artificial chromosome, and yeast artificial chromosome.
- 77. (Amended) A-<u>The</u> vector according to claim 76, wherein said bacteriophage is selected from the group consisting of λ gt10 and λ gt11 and phage display vectors.
- 78. (Amended) A-<u>The</u> vector according to claim 77, wherein said phage display vector is selected from vectors derived from pCOMB vectors.
- 79. (Amended) A-<u>The</u> vector according to claim 76-or-77, wherein said phage display vector is of the MCO group.

- 80. (Amended) A-The vector according to any one of claims claim 77-to-79, wherein said phage display vector is selected from the group consisting of MCO1, MCO3 and MCO6 vectors.
- 81. (Amended) A-<u>The</u> vector according to any one of claims claim 77-to-80, wherein said phage display vector is MCO3.
- 82. (Amended) A-<u>The</u> vector according to claim 75, wherein said vector is a mammalian expression vector.
- 83. (Amended) A-<u>The</u> vector according to claim 82, wherein said mammalian expression vector is pG1D102-MCO or pKN100-MCO.
- 84. (Amended) A host cell transformed with the vector according to any one of claims claim 75 to 83.
- 85. (Amended) A-<u>The</u> host cell according to claim 84, wherein said host cell is selected from the group consisting of *E. coli*, *Bacillus*, *Streptomyces*, *Pseudomonas*, *Salmonella*, and *Serratia*.
- 86. (Amended) A-The host cell according to claim 84, wherein said host cell is selected from the group consisting of yeast, fungi, plant, insect cells and mammalian cells.
- 87. (Amended) A-<u>The</u> host cell according to claim 86, wherein said mammalian cells are selected from the group consisting of CHO cell lines, COS cell lines, HeLa cells, L cells, murine 3T3 cells, c6 glioma cells and myeloma cell lines.
- 88. (Amended) A-<u>The</u> host cell according to claim 86 or claim 87, wherein said mammalian cells are CHO DG44 cells.
- 89. (Amended) A non-human vertebrate comprising a host cell according to any one of claimsclaim 84 to 88.
- 90. (Amended) A pharmaceutical composition comprising the polypeptide according to any one of claims 27 to 47, or a peptide fragment according to any one of claims 48 to 54, or an antibody or fragment thereof according to any one of claims 55 to 74, together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.

- 91. (Amended) A-The pharmaceutical composition according to claim 90, wherein said polypeptide is in a form selected from the group consisting of polypeptide/chelate, polypeptide/drug, polypeptide/prodrug, polypeptide/toxin, polypeptide/imaging marker, antibody/chelate, antibody/drug, antibody/prodrug, antibody/toxin and antibody/imaging marker.
- 92. (Amended) A-<u>The</u> pharmaceutical composition according to claim 91, wherein said chelate is selected from the group consisting of: ⁹⁰Y, ¹³¹I and ¹⁸⁸Re.
- 93. (Amended) A-<u>The</u> pharmaceutical composition according to claim 91, wherein said drug is a cytotoxic drug.
- 94. (Amended) A-The pharmaceutical composition according to claim 93, wherein said cytotoxic drug is selected from the group consisting of adriamycin, melphalan, cisplatin, taxol, fluorouricil, cyclophosphamide
- 95. (Amended) A-The pharmaceutical composition according to claim 91, wherein said prodrug is an antibody directed prodrug therapy or ADEPT.
- 96. (Amended) A-<u>The</u> pharmaceutical composition according to claim 91, wherein said toxin is selected from the group consisting of ricin, abrin, *Diptheria* toxin and *Pseudomonas* endotoxin (PE 40).
- 97. (Amended) A-<u>The</u> pharmaceutical composition according to claim 91, wherein said imaging marker is selected from the group consisting of ¹²⁵l, ¹³¹l, ¹²³l, ¹¹¹ln, ¹⁰⁵Rh, ¹⁵³Sm, ⁶⁷Cu, ⁶⁷Ga, ¹⁶⁶Ho, ¹⁷⁷Lu, ¹⁸⁶Re, ¹⁸⁸Re, and ^{99m}Tc.
- 98. (Amended) A-<u>The</u> pharmaceutical composition according to claim 91, wherein said imaging marker is gadolinium.
- 99. (Amended) A vaccine comprising a nucleic acid sequence according to any one of claimsclaim 1 to 26, or a fragment thereof, or a polypeptide according to any one of claims 27 to 47, or a peptide fragment according to any one of claims 48 to 54, or an antibody or fragment thereof according to any one of claims 55 to 74, together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.

- 100. (Amended) A-The vaccine according to claim 99, wherein said vaccine is an idiotypic vaccine.
- 101. (Amended) A-The vaccine according to claim 99-or claim 100, wherein said vaccine is formulated for administration via an oral, inhalation, topical or parenteral route.
- 102. (Amended) A method for inducing an immune response against disease in a vertebrate, comprising administering to said vertebrate an immunologically effective amount of the polypeptide-(_or peptide fragment thereof)__according to any one of claims described to any one of claims 48 to 54, or an antibody (or fragment thereof) according to any one of claims 55 to 74, or a pharmaceutical composition according to any one of claims 90 to 98, or a vaccine according to any one of claims 99 to 101.
- 103. (Amended) The method according to claim 102, wherein the polypeptide, peptide fragment, or antibody-(_or fragment thereof)-_is administered together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 104. (Amended) A method for the treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to said vertebrate a therapeutically effective amount of the polypeptide-{_, or peptide fragment thereof}-_, according to any one of claims claim 27 to 47, or the peptide fragment according to any one of claims 48 to 54, or an antibody (or fragment thereof) according to any one of claims 55 to 74, or a pharmaceutical composition according to any one of claims 90 to 98, or a vaccine according to any one of claims 99 to 101.
- 105. (Amended) The method according to any one of claims claim 102 to 104, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 106. (Amended) The method according to any one of claims claim 102 to 105, wherein the disease is cancer.
- 107. (Amended) The method according to claim 106, wherein the cancer is selected from the group consisting of carcinogenic tumourtumors; tumourtumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumourtumors, hepatic

cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumourtumors, such as sarcoma; and haemopoietic tumourtumors, such as B cell lymphoma.

- 108. (Amended) A diagnostic kit for the detection of polypeptides encoded by the p53 gene in vertebrates, said kit comprising the antibody-(_or fragment thereof)-_according to any one of claimsclaim 55 to 74, together with a diagnostically acceptable carrier and/or diluent.
- 109. (Amended) A-The diagnostic kit according to claim 108, wherein said kit comprises:
 - (a) a first container containing the antibody-{, or fragment thereof}-, wherein said antibody or fragment thereof has binding affinity to a p53 protein or a portion thereof in vertebrates, and wherein said antibody is obtained from a vertebrate host expressing an immune response against a naturally-occurring disease.according to any one of claims 55 to 74, and;
 - (b) a second container containing a conjugate comprising a binding partner of the antibody-(_or fragment thereof), together with a detectable label.
- 110. (Amended) A method for screening for a disease in a vertebrate comprising:
 - (a) contacting a sample from a vertebrate with a nucleic acid probe comprising a nucleic acid sequence according to any one of claimsclaim 1-to-26, or an oligonucleotide fragment thereof, and
 - (b) detecting <u>hybridisation hybridization</u> between the nucleic acid sample and the polynucleotide sequence.
- 111. (Amended) A-The method according to claim 110, wherein the oligonucleotide fragment is between about 10 to about 100 nucleotides in length.
- 112. (Amended) A-<u>The</u> method according to claim 110-or claim 111, wherein the oligonucleotide fragment is between about 15 to about 30 nucleotides in length.
- 113. (Amended) The method according to any one of claims claim 110 to 112, wherein hybridisation hybridization as compared to non-hybridisation hybridization is indicative of disease.

- 114. (Amended) The method according to any one of claims claim 110 to 113, wherein said disease is cancer.
- 115. (Amended) The method according to any one of claimsclaim 110 to 114, wherein hybridisation is conducted under low, moderate, or high stringency.
- 116. (Amended) The method according to any one of claims claim 110 to 115, wherein hybridisation is conducted under high stringency.
- 117. (Amended) A method for screening for a disease in a vertebrate comprising:
 - (a) contacting a sample from a vertebrate with the antibody-(_or fragment thereof)-, according to any one of claimsclaim 55 to 74, and
 - (b) detecting the presence of the antibody-(_or fragment thereof)-_bound to a p53 polypeptide.
- 118. (Amended) A-The method according to claim 117, wherein said disease is cancer.
- 119. (Amended) A method of gene therapy, wherein said method comprises:
 - (a) inserting a nucleic acid sequence according to any one of claimsclaim 1-to 26, or a vector according to any one of claimsclaim 75 to 83, into a host cell;
 - (b) expressing the nucleic acid sequence in the transformed cell.
- 121. (Amended) A process for preparing an antibody-{_or fragment thereof}-_having binding affinity to a p53 protein or a portion thereof in vertebrates, wherein said process comprises:
 - (a) isolating from a vertebrate a nucleic acid sequence according to any one of claimsclaim 1-to-26;
 - (b) cloning said nucleic acid sequence into a vector;
 - (c) constructing an antibody fragment library; and
 - (d) screening said library for clones expressing the antibody of interest.
- 122. (Amended) The process according to claim 121, wherein said antibody-(_or fragment thereof)-_has binding affinity to a p53 protein or a portion thereof in vertebrates.
- 125. (Amended) The process according to any one of claims claim 121-to-124, wherein the vector is a phage display vector.

- 127. (Amended) The process according to claim 125-or claim-126, wherein the vector is MCO1.
- 128. (Amended) A method of locating a nucleotide sequence encoding a polypeptide of an antibody-(_or fragment thereof)-_having binding affinity to a p53 protein or portion thereof in vertebrates, using the nucleic acid sequence according to any one of claimsclaim 1-to-26, or an oligonucleotide fragment thereof.
- 129. (Amended) The method according to claim 128, comprising:
 - (a) contacting a biological sample with a nucleic acid sequence according to comprising a polynucleotide sequence encoding a polypeptide of an antibody, or fragment thereof, wherein said antibody, or fragment thereof, has binding affinity to a p53 protein or a portion thereof in vertebrates, and wherein said nucleic acid sequence is obtained from a vertebrate host expressing an immune response against a naturally-occurring diseaseany one of claimsclaim 1 to 26, or an oligonucleotide fragment thereof; and
 - (b) identifying nucleotide sequences in the biological sample which hybridise hybridize to said nucleic acid sequence or oligonucleotide fragment.
- 130. (Amended)—— A-The method according to claim 129, wherein the oligonucleotide fragment is between about 10 to about 100 nucleotides in length.
- 131. (Amended) A-<u>The</u> method according to claim 129-or claim 130, wherein the oligonucleotide fragment is between about 15 to about 30 nucleotides in length.
- 132. (New) A pharmaceutical composition comprising a peptide fragment according to claim 48 together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 133. (New) A pharmaceutical composition comprising an antibody or fragment thereof according to claim 55 together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 134. (New) A vaccine comprising a polypeptide according to claim 27 together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.

- 135. (New) The vaccine according to claim 134, wherein said vaccine is an idiotypic vaccine.
- 136. (New) The vaccine according to claim 134, wherein said vaccine is formulated for administration via an oral, inhalation, topical or parenteral route.
- 137. (New) A vaccine comprising a peptide fragment according to claim 48 together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 138. (New) The vaccine according to claim 137, wherein said vaccine is an idiotypic vaccine.
- 139. (New) The vaccine according to claim 137, wherein said vaccine is formulated for administration via an oral, inhalation, topical or parenteral route.
- 140. (New) A vaccine comprising an antibody or fragment thereof according to claim 55, together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 141. (New) The vaccine according to claim 140, wherein said vaccine is an idiotypic vaccine.
- 142. (New) The vaccine according to claim 140, wherein said vaccine is formulated for administration via an oral, inhalation, topical or parenteral route.
- 143. (New) A method for inducing an immune response against disease in a vertebrate, comprising administering to said vertebrate an immunologically effective amount of the peptide fragment according to claim 48.
- 144. (New) The method according to claim 143, wherein the polypeptide, peptide fragment, or antibody, or fragment thereof, is administered together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 145. (New) A method for the treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to said vertebrate a therapeutically effective amount of the peptide fragment according to claim 48.
- 146. (New) The method according to claim 143, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.

- 147. (New) The method according to claim 143, wherein the disease is cancer.
- 148. (New) The method according to claim 147, wherein the cancer is selected from the group consisting of carcinogenic tumors; tumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumors, such as sarcoma; and haemopoietic tumors, such as B cell lymphoma.
- 149. (New) A method for inducing an immune response against disease in a vertebrate, comprising administering to said vertebrate an immunologically effective amount of the antibody, or fragment thereof, according to claim 55.
- 150. (New) The method according to claim 149, wherein the polypeptide, peptide fragment, or antibody, or fragment thereof, is administered together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 151. (New) A method for the treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to said vertebrate a therapeutically effective amount of the antibody, or fragment thereof, according to claim 55.
- 152. (New) The method according to claim 149, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 153. (New) The method according to claim 149, wherein the disease is cancer.
- 154. (New) The method according to claim 153, wherein the cancer is selected from the group consisting of carcinogenic tumors; tumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumors, such as sarcoma; and haemopoietic tumors, such as B cell lymphoma.
- 155. (New) A method for inducing an immune response against disease in a vertebrate, comprising administering to said vertebrate an immunologically effective amount of the pharmaceutical composition according to claim 90.

- 156. (New) The method according to claim 155, wherein the polypeptide, peptide fragment, or antibody, or fragment thereof, is administered together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 157. (New) A method for the treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to said vertebrate a therapeutically effective amount of the pharmaceutical composition according to claim 90.
- 158. (New) The method according to claim 155, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 159. (New) The method according to claim 155, wherein the disease is cancer.
- 160. (New) The method according to claim 159, wherein the cancer is selected from the group consisting of carcinogenic tumors; tumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumors, such as sarcoma; and haemopoietic tumors, such as B cell lymphoma.
- 161. (New) A method for inducing an immune response against disease in a vertebrate, comprising administering to said vertebrate an immunologically effective amount of the vaccine according to claim 99
- 162. (New) The method according to claim 161, wherein the polypeptide, peptide fragment, or antibody, or fragment thereof, is administered together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 163. (New) A method for the treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to said vertebrate a therapeutically effective amount of the vaccine according to claim 99.
- 164. (New) The method according to claim 161, wherein the disease is selected from the group consisting of cancer, rheumatoid arthritis and coronary heart disease.
- 165. (New) The method according to claim 161, wherein the disease is cancer.

- 166. (New) The method according to claim 165, wherein the cancer is selected from the group consisting of carcinogenic tumors; tumors of epithelial origin, such as colo-rectal cancer, breast cancer, lung cancer, head and neck tumors, hepatic cancer, pancreatic cancer, ovarian cancer, gastric cancer, brain cancer, bladder cancer, prostate cancer and urinary/genital tract cancer, oesophageal cancer; mesenchymal tumors, such as sarcoma; and haemopoietic tumors, such as B cell lymphoma.
- 167. (New) A method of gene therapy, wherein said method comprises:
 - (a) inserting a vector according to claim 75 into a host cell;
 - (b) expressing the nucleic acid sequence in the transformed cell.